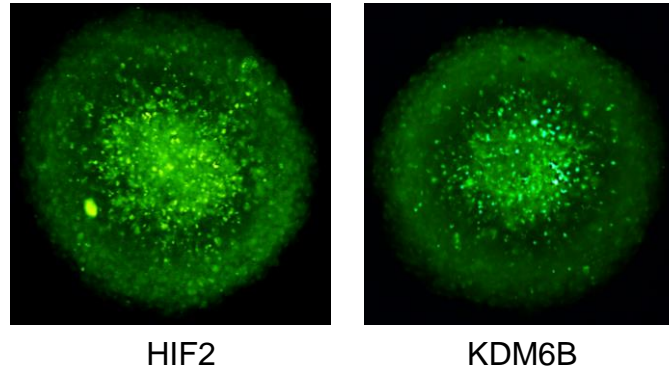


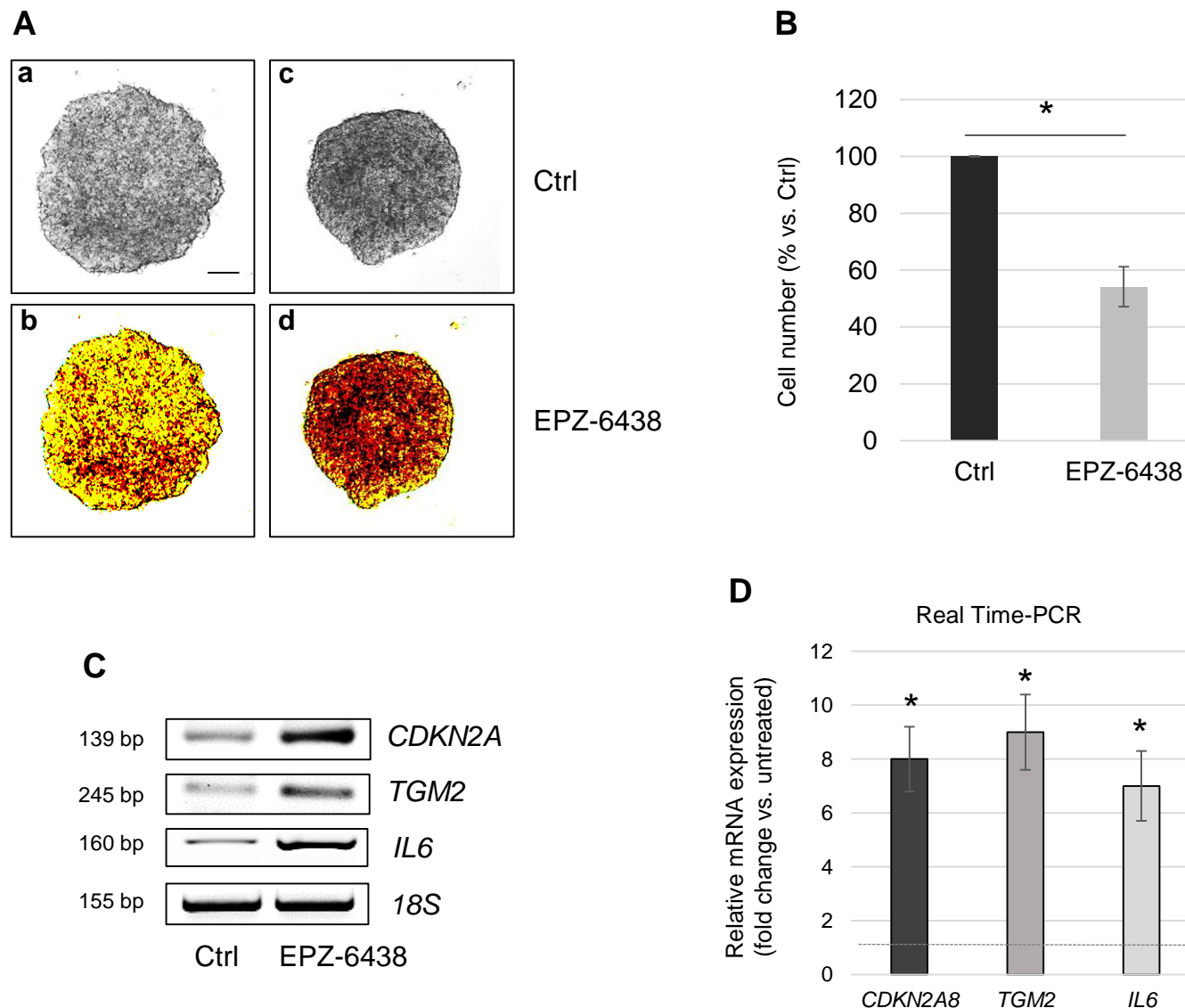
**Figure S1**

**a-f)** Representative images of immunostaining. Tumor biopsies from 6 patients diagnosed with BAP1<sup>+</sup> biphasic MPM were stained for H3K27me3. Images were captured at magnification x40 using light microscopy. Arrows indicate highly positive infiltrating inflammatory cells.



**Figure S2**

Representative images of immunofluorescence. MSTO-211H spheroids were stained for HIF2 and KDM6B. Images were captured using fluorescence microscopy (x40 magnification).



**Figure S3**

**A)** Representative phase contrast images (x40 magnification) (a,c) and relative pseudo-color images (b,d) of BR95 multicellular spheroids treated for 48 hours, with or without EPZ-6438. **B)** Bar graph shows the percentage of viability of BR95 cells cultured as monolayer (2D)  $\pm$  48 hours treatment with EPZ-6438. Each bar represents mean of three independent experiments  $\pm$  s.d., \* $p \leq 0.05$ . **C)** Representative RT-PCR and Real Time PCR **D)** analysis of *CDKN2A*, *TGM2*, and *IL6* mRNA expression in BR95 spheroids treated or not 48 hours with EPZ-6438. 18S rRNA was used as housekeeping gene, \* $p \leq 0.05$ .